

What is claimed is:

1. A method of assaying the activity of a fatty acid amide hydrolase comprising the steps of:

combining a sample suspected of containing a fatty acid amide hydrolase, with a labeled substrate of the fatty acid amide hydrolase, to form a reaction mixture;

incubating the reaction mixture under conditions sufficient to allow the fatty acid amide hydrolase to hydrolyze the labeled substrate, thereby forming at least one labeled hydrolysis product;

contacting the incubated reaction mixture with a selective binding material; wherein the selective binding material binds either the labeled substrate or a labeled hydrolysis product, but not both, thereby forming a bound labeled complex; separating the bound labeled complex from the incubated reaction mixture;

and

determining an amount of labeled substrate hydrolyzed, or labeled hydrolysis product formed, thereby indicating the fatty acid amide hydrolase activity of the sample.

- 2. The method of claim 1 wherein the sample comprises biological membranes, lipid bilayers, or micelles.
- 3. The method of claim 1 wherein the substrate is an endocannabinoid, a fatty acid ethanolamide, a fatty acid primary amide, an endocannabinoid analog, a fatty acid ethanolamide analog, or a fatty acid primary amide analog.
- 4. The method of claim 1 wherein the substrate is anandamide.

- 5. The method of claim 1 wherein the substrate is oleamide.
- 6. The method of claim 1 wherein the substrate is 2-arachidonoylglycerol.
- 7. The method of claim 1 wherein the substrate is labeled with a radioisotope.
- 8. The method of claim 7 wherein the radioisotope is ³H or ¹⁴C
- 9. The method of claim 1 wherein the substrate is labeled with a fluorescent label.
- 10. The method of claim 1 wherein the selective binding material comprises carbon.
- 11. The method of claim 10 wherein the selective binding material is activated charcoal.
- 12. The method of claim 11 wherein the activated charcoal comprises a filter.
- 13. The method of claim 1 wherein the selective binding material binds the labeled substrate but not the labeled product.
- 14. The method of claim 1 wherein the separating step comprises filtration, gravity settling or centrifugation.



- 15. The method of claim 1 wherein the determining step is performed via liquid scintillation counting or by measurement of fluorescence energy.
- 16. The method of claim 1 conducted in a multiwell plate.
- 17. The method of claim 1 comprising at least a portion of a high throughput screening program.
- 18. The method of claim 1 wherein the method is conducted in conjunction with a drug discovery effort.
- 19. A method of identifying a compound that modulates the activity of a fatty acid amide hydrolase comprising the steps of:

comparing the activity of a fatty acid amide hydrolase as assayed by the method of claim 1, in the presence and in the absence of a test compound added to the reaction mixture;

wherein a change in the activity of the fatty acid amide hydrolase indicates that the test compound modulates the activity of the fatty acid amide hydrolase.

- 20. The method of claim 19 wherein the test compound is selected from a library of compounds.
- 21. The method of claim 19 wherein the test compound inhibits the activity of the fatty acid amide hydrolase activity.



- 22. The method of claim 21 wherein said fatty acid amide hydrolase activity is inhibited at least about 5%.
- 23. The method of claim 21 wherein said fatty acid amide hydrolase activity is inhibited at least about 20%.
- 24. The method of claim 21 wherein said fatty acid amide hydrolase activity is inhibited at least about 50%.
- 25. The method of claim 21 wherein said fatty acid amide hydrolase activity is inhibited at least about 80%.
- 26. The method of claim 21 wherein said fatty acid amide hydrolase activity is inhibited at least about 95% or more.
- 27. The method of claim 21 wherein said test compound increases said fatty acid amide hydrolase activity.
- 28. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased at least about 5%.
- 29. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased at least about 30%.

- 30. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased at least about 50%.
- 31. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased at least about 70%.
- 32. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased at least about 100%.
- 33. The method of claim 27 wherein said fatty acid amide hydrolase activity is increased between about two-fold to about ten-fold.
- 34. The method of claim 19, which comprises the use of a multi-well plate.
- 35. The method of claim 19 conducted in a multiwell plate.
- 36. The method of claim 19 comprising at least a portion of a high throughput screening.
- 37. The method of claim 19 wherein the method is conducted in conjunction with a drug discovery effort.
- 38. The method of claim 1 wherein said fatty acid amide hydrolase is a mammalian fatty acid amide hydrolase.



- 39. The method of claim 38 wherein said fatty acid amide hydrolase is a porcine fatty acid amide hydrolase.
- 40. The method of claim 38 wherein said fatty acid amide hydrolase is a rodent fatty acid amide hydrolase.
- 41. The method of claim 38 wherein said fatty acid amide hydrolase is a murine fatty acid amide hydrolase.
- 42. The method of claim 41 wherein said fatty acid amide hydrolase is a rat fatty acid amide hydrolase.
- 43. The method of claim 41 wherein said fatty acid amide hydrolase is a mouse fatty acid amide hydrolase.
- 44. The method of claim 38 wherein said fatty acid amide hydrolase is a human fatty acid amide hydrolase.
- 45. A method for determining altered fatty acid amide hydrolase activity in a patient comprising:

obtaining a sample containing cells from the patient;

lysing the cells to form a cell lysate;

combining the cell lysate with a labeled substrate of fatty acid amide hydrolase, to form a reaction mixture;

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incubating the reaction mixture under conditions sufficient to allow a fatty acid amide hydrolase present in the cell lysate to hydrolyze the labeled substrate, thereby forming at least one labeled hydrolysis product;

contacting the incubated reaction mixture with a selective binding material; wherein the selective binding material binds either the labeled substrate or a labeled hydrolysis product, but not both, thereby forming a bound labeled complex;

separating the bound labeled complex from the incubated reaction mixture;

determining an amount of labeled substrate hydrolyzed, or labeled hydrolysis

product formed, thereby indicating the fatty acid amide hydrolase activity of the

sample; and

comparing the activity of the sample from the patient with the activity of a to a predetermined value for activity, to determine if the patient has altered fatty acid amide hydrolase activity relative to the predetermined value for activity.

- 46. The method of claim 45 wherein said patient is female.
- 47. The method of claim 46 wherein the female is pregnant or is seeking fertility treatment.
- 48. The method of claim 45 wherein the sample comprises blood, tissue or body fluid.
- 49. The method of claim 46 wherein the sample comprises lymphocytes.
- 50. The method of claim 45 wherein the cells are homogenized.



- 51. The method of claim 45 wherein the fatty acid amide hydrolase activity present in the cell lysate is partially or substantially purified from the sample.
- 52. The method of claim 45 wherein the predetermined determined value is from a control assay, a prior or subsequent sample from the patient, a sample from a normal individual, a sample from another patient, a standard FAAH, or a predetermined value.